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CLAIMS

- a domain capable of binding selectively a defined DNA sequence and a detecting domain capable of binding specifically a transactivator or a transrepressor or a transactivating or transrepressing complex characteristic of a physiological or physiopathological state.
- 2. Molecule according to claim 1,

 10 characterized in that the domain capable of binding selectively a defined DNA sequence is derived from a protein capable of interacting with the DNA.
- 3. Molecule according to claim 2, characterized in that the domain capable of binding selectively a defined DNA sequence is derived from a eukaryotic protein.
 - 4. Molecule according to claim 3, characterized in that the domain capable of binding selectively a defined DNA sequence is derived from the proteins p53, STAT or NFkB.
 - 5. Molecule according to claim 2, characterized in that the domain capable of binding selectively a defined DNA sequence is derived from a prokaryotic protein.
- 25 6. Molecule according to claim 5, characterized in that the prokary tic protein is a bacterial repressor.
 - 7. Molecule according to claim 6,

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characterized in that the domain capable of binding selectively a defined DNA sequence is derived from the tetR protein.

- 8. Molecule according to claim 6,
 5 characterized in that the domain capable of binding selectively a defined DNA sequence is derived from the Cro protein.
 - 9. Molecule according to one of claims 2 to 8, characterized in that the domain capable of binding selectively a defined DNA sequence comprises the domain for interacting with DNA of the said protein.
 - 10. Molecule according to one of claims 2 to 8, characterized in that the domain capable of binding selectively a defined DNA sequence consists of a complete protein.
 - 11. Molecule according to claim 10, characterized in that the domain capable of binding selectively a defined DNA sequence consists of the tetR protein.
- 20 12. Molecule according to claim 10, characterized in that the domain capable of binding selectively a defined DNA sequence consists of the Croprotein.
- 13. Molecule according to claim 1,

 25 characterized in that the domain capable of binding specifically the transactivator or transrepressor or the transactivating or transrepressing complex is an oligomerizing domain.

- 14. Molecule according to claim 13, characterized in that the oligomerizing domain is a leucine zipper, an SH3 or SH2 domain.
- 15. Molecule according to claim 13,

 5 characterized in that the oligomerizing domain capable of binding specifically the transactivator consists of the C-terminal part of the p53 protein.
 - 16. Molecule according to claim 15, characterized in that the oligomerizing domain consists of the C-terminal part of the p53 protein ranging from residues 320 to 393 (SEQ ID No. 3), 302-360 or 302-390.
- 17. Molecule according to claim 1,
 characterized in that the domain capable of binding
 specifically the transactivator or transrepressor or
 15 the transactivating or transrepressing complex is a
 synthetic or natural domain known to interact with the
 said transactivator or transrepressor or
 transactivating or transrepressing complex.
- characterized in that the domain capable of binding specifically the transactivator or transrepressor or the transactivating or transrepressing complex is an antibody or an antibody fragment or derivative directed against the transactivator or transrepressor or transactivating or transrepressor or
 - 19. Molecule according to claim 18, charact rized in that the domain capable of binding specifically the transactivator or the transactivating

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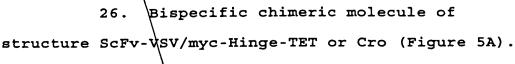
complex consists of a Fab or F(ab)'2 fragment of antibodies or a VH or VL region of an antibody.

- 20. Melecule according to claim 18, characterized in that the domain capable of binding specifically the transactivator or the transactivating complex consists of a single-chain antibody (ScFv) comprising a VH region linked to a VL region by an arm.
- 21. Molecule according to claim 1, characterized in that the DNA-binding domain and the transactivator binding domain are linked to each other through an arm.
- 20. Molecule according to claim 21, characterized in that the arm consists of a peptide comprising 5 to 30 amino acids and, preferably, 5 to 20 amino acids.
- 23. Molecule according to claim 22, characterized in that the arm is chosen from the peptides of sequence SEQ ID No. 5 or SEQ ID No. 6.
- 24. Molecule according to one of the
 20 preceding claims, characterized in that the DNA-binding domain is situated at the N-terminal position and the transactivator-binding domain is situated at the C-terminal position.
- 25. Molecule according to one of claims 1 to
 25 23, characterized in that the DNA-binding domain is
 situated at the C-terminal position and the
 transactivator-binding domain is situated at the
 N-terminal position.

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27. Bispecific chimeric molecule of structure ScFv-Hinge-TET or Cro (Figure 5B).

28. Bispecific chimeric molecule of structure ScFv-TET or Cro (Figure 5C).

29. Bispecific chimeric molecule of structure TET or Cro-ScFv (Figure 5D).

30. Bispecific chimeric molecule of to structure TET or Cro-Hinge-ScFv (Figure 5E).

31. Bispecific chimeric molecule of structure Oligom-VSV/myc-Hinge-TET or Cro (Figure 5A), Oligom-Hinge-TET or Cro (Figure 5B) or Oligom-TET or Cro (Figure 5C).

32. Nucleic acid sequence encoding a Clarm / chimeric molecule according to one of claims 1 to 31.

33. Nucleic acid sequence according to claim 32, characterized in that it is a DNA sequence.

34. Nucleic acid sequence according to claim 32 or 33, characterized in that it is part of a vector.

35. Conditional system for the expression of genes comprising:

- a chimeric molecule as defined in claims 1

- an expression cassette comprising a regulatory s quence, a minimal transcriptional promoter and the said gene.

36. Conditional system according to claim

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35, characterized in that the DNA-binding domain of the chimeric molecule is represented by all or part of TetR and the regulatory sequence comprises the sequence SEQ ID No. 1 or a derivative thereof, optionally repeated several times.

- 37. Conditional system according to claim
 35, characterized in that the DNA-binding domain of the chimeric molecule is represented by all or part of Cro and the regulatory sequence comprises the sequence SEQ ID No. 2 or a derivative thereof, optionally repeated several times.
- 38. Conditional system according to one of claims 35 to 37, characterized in that the minimal promoter comprises an INR or TATA box.
- 39. Conditional system according to claim
 38, characterized in that the minimal promoter is
 derived from the promoter of the thymidine kinase gene.
- 40. Conditional system according to claim 39, characterized in that the minimal promoter is composed of nucleotides -37 to +19 of the promoter of the thymidine kinase gene.
 - 41. Vector comprising:
- a nucleic acid sequence encoding a chimeric molecule according to one of claims 1 to 31, and
- an expression cassette comprising a regulatory sequence, a minimal transcriptional promoter and a coding sequence of interest.
 - 42. Vector according to claim 41,

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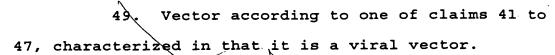
characterized in that the minimal transcriptional promoter is defined according to claims 38 to 40.

- 43. Vector according to claim 41, characterized in that the DNA-binding domain of the chimeric molecule is represented by all or part of TetR and the regulatory sequence comprises the sequence SEQ ID No. 1 or a derivative thereof, optionally repeated several times.
- 44. Vector according to claim 41,

 10 characterized in that the DNA-binding domain of the chimeric molecule is represented by all or part of Cro and the regulatory sequence comprises the sequence SEQ ID No. 2 or a derivative thereof, optionally repeated several times.
 - 45. Vector according to one of claims 41 to
 44, characterized in that the coding sequence of

 interest is a DNA sequence encoding a therapeutic
 product.
- 46. Vector according to claim 45,

 20 characterized in that the therapeutic product is a toxic polypeptide or peptide.
 - 47. Vector according to claim 46, characterized in that the toxic therapeutic product is chosen from diphtheria toxin, pseudomonas toxin, ricin A, thymidine kinase, cytosine deaminase, protein Grb3-3, or ScFv Y28.
 - 48. Vector according to one of claims 41 to 47, characterized in that it is a plasmid vector.



- 50. Vector according to claim 49, characterized in that it is a defective recombinant adenovirus.
- 51. Vector according to claim 49, characterized in that it is a defective recombinant retrovirus.
- 52. Pharmaceutical composition comprising at least one vector according to one of claims 41 to 51.
- 53. Nucleic acid comprising the sequence SEQ ID No. 4.
- 54. Molecule according to claim 1, characterized in that the transactivator characteristic of a physiological or physiopathological state is a protein of viral, parasitic, mycobacterial or cellular origin having a transcriptional transactivating activity.
- 55. Molecule according to claim 54,

 20 characterized in that the transactivator is a viral protein chosen from the HIV virus Tat protein, the papilloma virus E6/E7 proteins and the Epstein-Barr virus EBNA protein.
- 56. Molecule according to claim 54,

 25 characterized in that the transactivator is a cellular protein, preferably the mutated or wild-type p53 protein.
 - 57. Molecule according to claim 1,

characterized in that the transactivator or transactivating complex characteristic of a physiological or physiopathological state is a protein appearing in an infected or hyperproliferative cell.

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